

WHAT IS CLAIMED IS:

1. A method for identifying rare events in a biological sample, comprising:
 3. obtaining a source of cells;
 4. contacting the source with a binding agent specific for a cell specific marker associated with a rare event wherein the binding agent is bound to a magnetic bead and wherein the binding agent binds to cells in the source expressing the cell specific marker;
 9. separating cells bound by the binding agent from the source thereby obtaining a sub-population of cells enriched for the cell specific marker associated with the rare event;
 12. placing the enriched sample on a substrate;
 13. automatically scanning the substrate at a plurality of coordinates;
 15. automatically obtaining a plurality of images at locations on the substrate that comprise the enriched sample; and
 18. processing the plurality of image to identify the rare event.
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21. 2. The method according to claim 1, wherein the binding agent is an antibody.
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24. 3. The method according to claim 1, wherein the sub-population is enriched for carcinoma cells.
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27. 4. The method of claim 1, wherein the separating is done by positive selection.
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30 5. The method of claim 1, wherein the separating is done by
31 negative selection.

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33 6. The method of claim 2, wherein the antibody is monoclonal
34 or polyclonal.

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36 7. The method of claim 2, wherein the antibody recognizes an
37 epithelial marker.

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39 8. The method of claim 2, wherein the antibody is selected
40 to avoid cross reactivity with the beads.

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42 9. The method of claim 3, wherein the carcinoma cells are
43 from peripheral blood.

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45 10. The method of claim 1, further comprising:

46 (a) automatically identifying a coordinate of the rare event;
47 and

48 (b) automatically acquiring an image of the rare event, at the
49 location coordinates.

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51 11. The method of claim 1, wherein the rare event is detected
52 by immunohistochemistry.

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54 12. The method of claim 1, wherein the rare event is detected
55 by in situ hybridization.

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57 13. The method of claim 1, wherein the rare event is detected
58 by a stain.

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60 14. The method of claim 13, wherein the stain is a nucleic
61 acid dye selected from the group consisting of hematoxylin,

62 Giemsa stain, methyl green, Nuclear Fast-Red, Hoechst 33342,
63 Hoechst 33258, thiazole orange, DAPI, ethidium bromide,
64 propidium iodide, TOTO, YOYO-1, SYTOX Blue, SYTOX Green, 7-
65 Aminoactinomycin, 9-Amino-6-chloro-2-methoxyacridine, and
66 acridine homodimer.

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68 15. The method of claim 13, wherein the rare event is stained
69 with a cytoplasmic dye such as eosin or Kleihauer-Betke
70 cytochemical stain or a combination thereof.

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72 16. The method of claim 1, wherein the cell specific marker
73 is detected by a nuclear stain and counterstain.

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75 17. The method of claim 1, wherein the cell specific marker
76 is detected by immunohistochemistry, in situ hybridization,
77 staining or a combination thereof.

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79 18. The method of claim 1, wherein the image is a digital
80 image.

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